entel

Glu,

wherein at least one of the following conditions (I) - (V) is true:

- I) at least one of  $X_4$ ,  $X_5$ ,  $X_6$ , Thr, Lys, and Arg is independently substituted with a non-natural or unusual amino acid,
  - II) the polypeptide is cyclized,
  - III) the polypeptide is stabilized,
  - IV) the aminoterminal amino acid residue is acylated, or
- V) the carboxyterminal amino acid residue is amidated, if the polypeptide is not cyclized, said sequence SEO ID NO:19 corresponding essentially to the C-terminal of said polypeptide, said polypeptide having at least one of the following properties:
- a) induces inhibition of spontaneous IL-8 production by human monocytes,
  - b) induces inhibition of IL-1 $\beta$  induced IL-8 production by human peripheral blood mononuclear cells (PBMC),
- c) induces production of interleukin-1 receptor antagonistic protein (IRAP) by human monocytes,
- d) induces chemotactic migration of CD8+ human Tlymphocytes in vitro,
- e) desensitizes human CD8+ T cells resulting in an unresponsiveness towards rhIL-10,  $\begin{picture}(20,0) \put(0,0){\line(0,0){100}} \put(0,0){\lin$
- f) . suppresses the chemotact c response of CD4+ T human lymphocytes towards IL-8,  $% \left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\right\} =\frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{ \frac{1}{2}\left\{$
- g) suppresses the chemotactic response of human monocytes towards MCAF/MCP-1,
- h) inhibits class II MHC molecule expression on human monocytes stimulated by IFN- $\gamma$ ,
- i) induces the production of IL-4 by cultured normal human CD4+ T cells,
- j) reduces [the] TNF $\alpha$  production in human mixed leukocyte reaction, or
- k) downregulates TNF $\alpha$  and IL-8 production in a rabbit model of bile acid induced acute pancreat tis and reduces

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neutrophi, infiltration in the lungs of the treated rabbits.

## Please add the following new claims:

- --65. The polypeptide of claim 18 where SEQ ID NO:19 is the C-terminal of said polypeptide and the polypeptide is not cyclized.
- 66. The polypeptide of claim 65 which has a length of up to about 20 amino acids.
- 67. The polypeptide of claim 66 whose length does not exceed 10 amino acids.
- 68. The polypeptide of claim 18, said polypeptide being selected from the group consisting of polypeptides identical to SEQ ID NO:19, SEQ ID NO:20, SEQ ID NO:21, and SEQ ID NO:22, except that at least one of conditions (I)-(V) applies.
- 69. The method of claim 49 which is a method of treating a disease.
- 70. The method of claim 69 where the disease involves proinflammatory activities.
- 71. The method of claim 69 where the disease is one inhibited by IL-10.
- 72. The method of claim 69 where the disease is one caused or aggravated by IL-8, MCAF or IL-1.--

## REMARKS

## 1. General Matters

- 1.1. As a result of this amendment, claims 18-72 are pending.
- 1.2. The Examiner is thanked for reconsidering the group restriction. If a generic claim is allowed, or is allowable in substance, the species restriction should be reconsidered, pursuant to MPEP \$809.02(c)(B) and 809.02(e).
  - 1.3. A §120 reference to PCT/DK97/00021 has been added.
  - 1.4. The "Figure legends" have been retitled and relocated.
  - 1.5. An abstract has been added.

## 2. Product of Nature Issues